**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicants: BONFANTI, Jean-François et al.

Serial No.: Not Assigned Art Unit: TBA

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For : 5- OR 6-SUBSTITUTED BENZIMIDAZOLE DERIVATIVES AS INHIBITORS OF RESPIRATORY SYNCYTIAL VIRUS REPLICATION

Mail Stop: PCT
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**PRELIMINARY AMENDMENT "A"**

Dear Sir:

Prior to examination and calculation of fees due, please amend the above-identified application as follows.

☒ Amendments to the Specification begin on page 2 of this paper.

☒ Amendments to the Claims are reflected in the listing of the claims which begins on page 3 of this paper.

☐ Amendments to the Drawings begin on page of this paper and include an attached replacement sheet.

☒ Remarks begin on page 18 of this paper.

**AMENDMENTS TO SPECIFICATION**

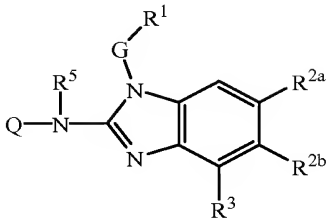
Page 1, between the Title and line 4, please insert the following new paragraph:

--Cross Reference to Related Applications

This application is the national stage of PCT Application No. PCT/EP2004/053618, filed December 20, 2004, which application claims priority from European Patent Application No. 03104806.9, filed 18 December 2003 and US provisional Application No. 60/566867, filed 30 April 2004, the entire disclosures of which are hereby incorporated in their entirely.--

Listing Claims

1. (Original) A compound having the formula

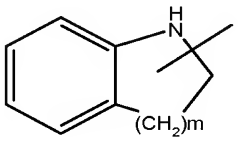


a prodrug, *N*-oxide, addition salt, quaternary amine, metal complex or stereochemically isomeric form thereof wherein

Q is $Ar^2$, $R^{6a}$, pyrrolidinyl substituted with $R^6$, piperidinyl substituted with $R^6$ or homopiperidinyl substituted with $R^6$;

G is a direct bond or $C_{1-10}$alkanediyl optionally substituted with one or more substituents individually selected from the group consisting of hydroxy, $C_{1-6}$alkyloxy, $Ar^1C_{1-6}$alkyloxy, $C_{1-6}$alkylthio, $Ar^1C_{1-6}$alkylthio, $HO(-CH_2-CH_2-O)_n-$, $C_{1-6}$alkyloxy$(-CH_2-CH_2-O)_n-$ and $Ar^1C_{1-6}$alkyloxy$(-CH_2-CH_2-O)_n-$;

$R^1$ is $Ar^1$ or a monocyclic or bicyclic heterocycle being selected from piperidinyl, piperazinyl, pyridyl, pyrazinyl, pyridazinyl, pyrimidinyl, furanyl, tetrahydrofuranyl, thienyl, pyrrolyl, thiazolyl, oxazolyl, imidazolyl, isothiazolyl, pyrazolyl, isoxazolyl, oxadiazolyl, quinolinyl, quinoxalinyl, benzofuranyl, benzothienyl, benzimidazolyl, benzoxazolyl, benzthiazolyl, pyridopyridyl, naphthiridinyl, 1*H*-imidazo[4,5-b]pyridinyl, 3*H*-imidazo[4,5-b]pyridinyl, imidazo[1,2-a]-pyridinyl, 2,3-dihydro-1,4-dioxino[2,3-b]pyridyl or a radical of formula



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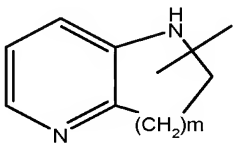

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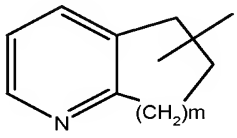

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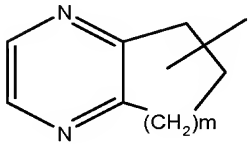

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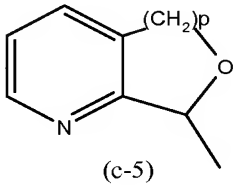

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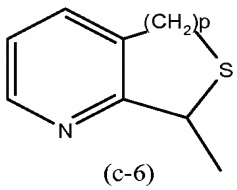

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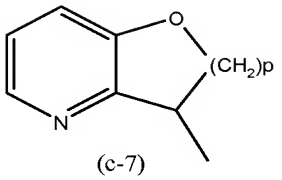


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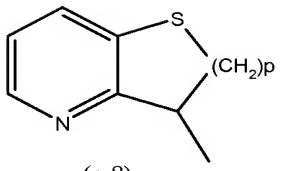



(c-8)

;

wherein each of said monocyclic or bicyclic heterocycles may optionally be substituted with 1 or where possible more, such as 2, 3, 4 or 5, substituents individually selected from the group of substituents consisting of halo, hydroxy, amino, cyano, carboxyl, $C_{1-6}$alkyl, $C_{1-6}$alkyloxy, $C_{1-6}$alkylthio, $C_{1-6}$alkyloxy$C_{1-6}$alkyl, $Ar^1$, $Ar^1C_{1-6}$alkyl, $Ar^1C_{1-6}$alkyloxy, hydroxy$C_{1-6}$alkyl, mono-or di($C_{1-6}$alkyl)amino, mono-or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, polyhalo$C_{1-6}$alkyl, $C_{1-6}$alkylcarbonylamino, $C_{1-6}$alkyl-$SO_2$-$NR^{4a}$-, $Ar^1$-$SO_2$-$NR^{4a}$-, $C_{1-6}$alkyloxycarbonyl, -C(=O)-$NR^{4a}R^{4b}$, HO(-$CH_2$-$CH_2$-O)$_n$-, halo(-$CH_2$-$CH_2$-O)$_n$-, $C_{1-6}$alkyloxy(-$CH_2$-$CH_2$-O)$_n$-, $Ar^1C_{1-6}$alkyloxy(-$CH_2$-$CH_2$-O)$_n$- and mono-and di($C_{1-6}$alkyl)amino(-$CH_2$-$CH_2$-O)$_n$-;

one of $R^{2a}$ and $R^{2b}$ is cyano$C_{1-6}$alkyl, cyano$C_{2-6}$alkenyl, $Ar^3C_{1-6}$alkyl, ($Ar^3$)(OH)$C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, N($R^{8a}R^{8b}$)$C_{1-6}$alkyl, $Ar^3C_{2-6}$alkenyl, Het-$C_{2-6}$alkenyl, $Ar^3$amino$C_{1-6}$alkyl, Het-amino$C_{1-6}$alkyl, Het-$C_{1-6}$alkylamino-$C_{1-6}$alkyl, $Ar^3$thio$C_{1-6}$alkyl, Het-thio$C_{1-6}$alkyl, $Ar^3$sulfonyl$C_{1-6}$alkyl, Het-sulfonyl-$C_{1-6}$alkyl, $Ar^3$aminocarbonyl, Het-aminocarbonyl, $Ar^3$($CH_2$)$_n$aminocarbonyl, Het-($CH_2$)$_n$aminocarbonyl, $Ar^3$carbonylamino, Het-carbonylamino, $Ar^3$($CH_2$)$_n$carbonylamino, Het-($CH_2$)$_n$carbonylamino, $Ar^3$($CH_2$)$_n$amino; and the other one of $R^{2a}$ and $R^{2b}$ is hydrogen;

in case $R^{2a}$ is hydrogen, then $R^3$ is hydrogen;

in case $R^{2b}$ is hydrogen, then $R^3$ is hydrogen or $C_{1-6}$alkyl;

$R^{4a}$ and $R^{4b}$ can be the same or can be different relative to one another, and are each independently hydrogen or $C_{1-6}$alkyl; or

$R^{4a}$ and $R^{4b}$ taken together may form a bivalent radical of formula -($CH_2$)$_s$- wherein s is 4 or 5;

$R^5$ is hydrogen or $C_{1-6}$alkyl;

$R^6$ is hydrogen or $C_{1-6}$alkyl optionally substituted with one or more substituents each independently selected from the group consisting of trifluoromethyl, $NR^{7a}R^{7b}$, $C_{3-7}$cycloalkyl, $Ar^2$, hydroxy, $C_{1-4}$alkoxy, $C_{1-4}$alkylthio, $Ar^2$-oxy-, $Ar^2$-thio-, $Ar^2$($CH_2$)$_n$oxy, $Ar^2$($CH_2$)$_n$thio, hydroxycarbonyl, aminocarbonyl, $C_{1-4}$alkyl-carbonyl, $Ar^2$carbonyl, $C_{1-4}$alkoxycarbonyl, $Ar^2$($CH_2$)$_n$carbonyl, amino-carbonyloxy, $C_{1-4}$alkylcarbonyloxy, $Ar^2$carbonyloxy, $Ar^2$($CH_2$)$_n$carbonyloxy, $C_{1-4}$alkoxycarbonyl($CH_2$)$_n$oxy, mono- or di($C_{1-4}$alkyl)aminocarbonyl, mono- or di($C_{1-4}$alkyl)aminocarbonyloxy, aminosulfonyl, mono- or di($C_{1-4}$alkyl)amino-sulfonyl or a heterocycle selected from the group consisting of pyrrolidinyl,

pyrrolyl, dihydropyrrolyl, imidazolyl, triazolyl, piperidinyl, homopiperidinyl, piperazinyl, pyridyl and tetrahydropyridyl, wherein each of said heterocycle may optionally be substituted with oxo or $C_{1-6}$alkyl;

$R^{6a}$ is $C_{1-6}$alkyl substituted with one or more substituents each independently selected from the group consisting of trifluoromethyl, $NR^{7a}R^{7b}$, $C_{3-7}$cycloalkyl, $Ar^2$, hydroxy, $C_{1-4}$alkoxy, $C_{1-4}$alkylthio, $Ar^2$-oxy-, $Ar^2$-thio-, $Ar^2(CH_2)_n$oxy, $Ar^2(CH_2)_n$thio, hydroxycarbonyl, aminocarbonyl, $C_{1-4}$alkylcarbonyl, $Ar^2$carbonyl, $C_{1-4}$alkoxycarbonyl, $Ar^2(CH_2)_n$carbonyl, aminocarbonyloxy, $C_{1-4}$alkylcarbonyl-oxy, $Ar^2$carbonyloxy, $Ar^2(CH_2)_n$carbonyloxy, $C_{1-4}$alkoxycarbonyl$(CH_2)_n$oxy, mono- or di($C_{1-4}$alkyl)aminocarbonyl, mono- or di($C_{1-4}$alkyl)aminocarbonyloxy, aminosulfonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl or a heterocycle selected from the group consisting of pyrrolidinyl, pyrrolyl, dihydropyrrolyl, imidazolyl, triazolyl, piperidinyl, homopiperidinyl, piperazinyl, pyridyl and tetrahydro-pyridyl, wherein each of said heterocycle may optionally be substituted with oxo or $C_{1-6}$alkyl;

$R^{7a}$ is hydrogen, $C_{1-6}$alkyl, formyl or $C_{1-6}$alkylcarbonyl;

$R^{7b}$ is hydrogen, $C_{1-6}$alkyl, formyl or $C_{1-6}$alkylcarbonyl;

$R^{8a}$ is $Ar^3$, $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, $C_{1-6}$alkoxy$C_{1-6}$alkyl, cyano$C_{1-6}$alkyl, amino$C_{1-6}$alkyl, mono-or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, aminocarbonyl-$C_{1-6}$-alkyl, carboxyl-$C_{1-6}$-alkyl;

$R^{8b}$ is $Ar^3$, $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, $C_{1-6}$alkoxy$C_{1-6}$alkyl, cyano$C_{1-6}$alkyl, amino$C_{1-6}$alkyl, mono-or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl;

each n independently is 1, 2, 3 or 4;

each m independently is 1 or 2;

each p independently is 1 or 2;

$Ar^1$ is phenyl or phenyl substituted with 1 or more, such as 2, 3 or 4, substituents selected from halo, hydroxy, $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, polyhalo$C_{1-6}$alkyl, and $C_{1-6}$alkyloxy;

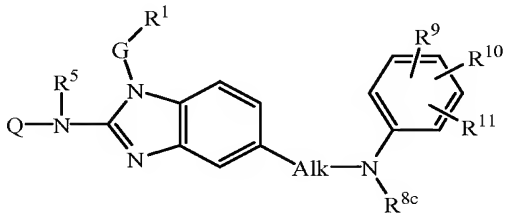
$Ar^2$ is phenyl or phenyl substituted with 1 or more, such as 2, 3 or 4, substituents selected from the group consisting of halo, hydroxy, amino, cyano, $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, polyhalo$C_{1-6}$alkyl, amino$C_{1-6}$alkyl, $C_{1-6}$alkyloxy, amino-sulfonyl, aminocarbonyl, hydroxycarbonyl, $C_{1-4}$alkylcarbonyl, mono- or di($C_{1-4}$alkyl)amino, mono- or di($C_{1-4}$alkyl)aminocarbonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl, mono- or di($C_{1-4}$alkyl)amino$C_{1-6}$alkyl and $C_{1-4}$alkoxycarbonyl;

$Ar^3$ is phenyl, naphthalenyl, 1,2,3,4-tetrahydro-naphthalenyl or indanyl, wherein said phenyl, naphtyl, 1,2,3,4-tetrahydro-naphthalenyl or indanyl may optionally and each individually be substituted with one or more, such as 2, 3 or 4, substituents

selected from the group consisting of halo, hydroxy, mercapto, amino, cyano, $C_{1-6}$alkyl, $C_{2-6}$alkenyl, $C_{2-6}$alkynyl, $Ar^1$, hydroxy$C_{1-6}$alkyl, polyhalo$C_{1-6}$alkyl, amino$C_{1-6}$alkyl, cyano$C_{1-6}$alkyl, aminocarbonyl, $C_{1-6}$alkyloxy, $C_{1-6}$alkylthio, $Ar^1$-oxy, $Ar^1$-thio, $Ar^1$-amino, aminosulfonyl, aminocarbonyl-$C_{1-6}$alkyl, hydroxycarbonyl-$C_{1-6}$alkyl, hydroxycarbonyl, $C_{1-4}$alkylcarbonyl, mono- or di($C_{1-4}$alkyl)amino, mono- or di($C_{1-4}$alkyl)aminocarbonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl, mono- or di($C_{1-4}$alkyl)amino$C_{1-6}$alkyl, $C_{1-4}$alkylcarbonylamino and $C_{1-4}$alkoxycarbonyl;

Het is a heterocycle being selected from tetrahydrofuranyl, tetrahydrothienyl, dioxanyl, dioxolanyl, pyrrolidinyl, pyrrolidinonyl, furanyl, thienyl, pyrrolyl, thiazolyl, oxazolyl, imidazolyl, isothiazolyl, pyrazolyl, isoxazolyl, oxadiazolyl, thiadiazolyl, piperidinyl, homopiperidinyl, piperazinyl, morpholinyl, pyridyl, pyrazinyl, pyridazinyl, pyrimidinyl, tetrahydroquinolinyl, quinolinyl, isoquinolinyl, benzodioxanyl, benzodioxolyl, indolinyl, indolyl, each of said heterocycle may optionally be substituted with oxo, amino, $Ar^1$, $C_{1-4}$alkyl, amino$C_{1-4}$alkyl, hydroxy$C_{1-6}$alkyl, $Ar^1C_{1-4}$alkyl, mono- or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, mono- or di($C_{1-6}$alkyl)amino, or with two $C_{1-4}$alkyl radicals.

2. (Original) A compound according to claim 1 wherein the compound has the formula (I-a-1):



(I-a-1)

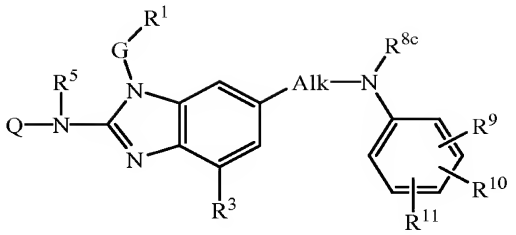
wherein Q, $R^5$, G and $R^1$ are as claimed in claim 1; and

Alk is $C_{1-6}$alkanediyl;

$R^{8c}$ has the same meanings of $R^{8a}$, as claimed in claim 1, and also may be hydrogen;

$R^9$, $R^{10}$, $R^{11}$ independently from one another have the same meanings as the substituents on $Ar^3$ as claimed in claim 1.

3. (Original) A compound according to claim 1 wherein the compound has the formula (I-b-1):



(I-b-1)

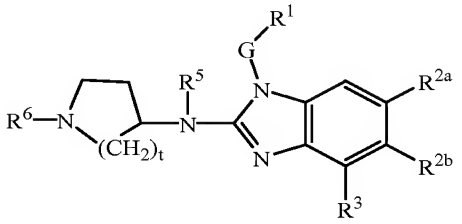
wherein Q, $R^5$, G and $R^1$ are as claimed in claim 1; and

Alk is $C_{1-6}$alkanediyl;

$R^{8c}$ has the same meanings of $R^{8a}$, as claimed in claim 1, and also may be hydrogen;

$R^9$, $R^{10}$, $R^{11}$ independently from one another have the same meanings as the substituents on $Ar^3$ as claimed in claim 1.

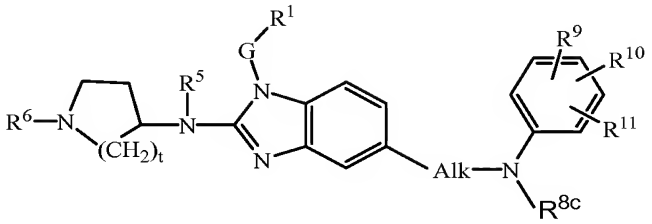
4. (Original) A compound according to claim 1 wherein the compound has the formula (I-c):



(I-c)

wherein t, G, $R^1$, $R^{2a}$, $R^{2b}$, $R^3$, $R^5$ and $R^6$ are as claimed in claim 1.

5. (Original) A compound according to claim 1 wherein the compound has the formula (I-d-2):



(I-d-2)

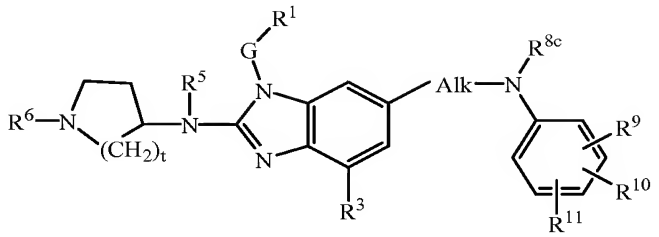
wherein t, $R^5$, $R^6$, G and $R^1$ are as claimed in claim 1; and

Alk is $C_{1-6}$alkanediyl;

$R^{8c}$ has the same meanings of $R^{8a}$, as claimed in claim 1, and also may be hydrogen;

$R^9$, $R^{10}$, $R^{11}$ independently from one another have the same meanings as the substituents on $Ar^3$ as claimed in claim 1.

6. (Original) A compound according to claim 1 wherein the compound has the formula (I-e-2):



(I-e-2)

wherein t, $R^5$, $R^6$, G and $R^1$ are as claimed in claim 1; and
Alk is $C_{1-6}$alkanediyl;
$R^{8c}$ has the same meanings of $R^{8a}$, as claimed in claim 1, and also may be hydrogen;
$R^9$, $R^{10}$, $R^{11}$ independently from one another have the same meanings as the substituents on $Ar^3$ as claimed in claim 1.

7. (Currently Amended) A compound according to claim 4~~any of claims 4 to 6~~ wherein t is 2.

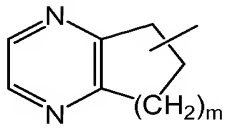
8. (Currently Amended) A compound according to claim 1~~any of claims 1 – 7~~, wherein G is $C_{1-10}$alkanediyl.

9. (Currently Amended) A compound according to claim 1~~in any of claims 1 – 7~~, wherein G is methylene.

10. (Currently Amended) A compound according to claim 1~~any of claims 1 – 9~~, wherein $R^1$ is pyridyl optionally substituted with 1 or 2 substituents independently selected from the group consisting of halo, hydroxy, amino, cyano, carboxyl, $C_{1-6}$alkyl, $C_{1-6}$alkyloxy, $C_{1-6}$alkylthio, $C_{1-6}$alkyloxy$C_{1-6}$alkyl, $Ar^1$, $Ar^1C_{1-6}$alkyl, $Ar^1C_{1-6}$alkyloxy, hydroxy$C_{1-6}$alkyl, mono-or di($C_{1-6}$alkyl)amino, mono-or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, polyhalo$C_{1-6}$alkyl, $C_{1-6}$alkylcarbonylamino, $C_{1-6}$alkyl-$SO_2$-$NR^{4a}$-, $Ar^1$-$SO_2$-$NR^{4a}$-, $C_{1-6}$alkyloxycarbonyl, -C(=O)-$NR^{4a}R^{4b}$, HO(-$CH_2$-$CH_2$-O)$_n$-, halo(-$CH_2$-$CH_2$-O)$_n$-, $C_{1-6}$alkyloxy(-$CH_2$-$CH_2$-O)$_n$-, $Ar^1C_{1-6}$alkyloxy(-$CH_2$-$CH_2$-O)$_n$- and mono-or di($C_{1-6}$alkyl)amino(-$CH_2$-$CH_2$-O)$_n$-.

11. (Currently Amended) A compound according to claim 1~~any of claims 1 – 9~~, wherein $R^1$ is pyridyl substituted with 1 or 2 substituents independently selected from the group consisting of hydroxy and $C_{1-6}$alkyl.

12. (Currently Amended) A compound according to claim 1~~any of claims 1 – 9~~, wherein $R^1$ is $Ar^1$, quinolinyl, benzimidazolyl, a radical of formula



(c-4)

or pyrazinyl; wherein each of the radicals $Ar^1$, quinolinyl, benzimidazolyl, (c-4), or pyrazinyl may optionally be substituted with the substitutents of said radicals as claimed in claim1.

13. (Currently Amended) A compound according to claim 1~~any of claims 1 – 9~~, wherein $R^1$ is phenyl optionally substituted with one, two or three radicals selected from the group consisting of halo, hydroxy, $C_{1-6}$alkyl, $C_{1-6}$alkyloxy; quinolinyl; a radical (c-4) wherein m is 2, optionally substituted with up to two radicals selected from $C_{1-6}$alkyl; benzimidazolyl optionally substituted with $C_{1-6}$alkyl; pyrazinyl optionally substituted with up to three radicals selected from $C_{1-6}$alkyl.

14. (Currently Amended) A compound according to claim 1~~any of claims 1 – 13~~, wherein one of $R^{2a}$ and $R^{3a}$ is selected from cyano$C_{1-6}$alkyl, cyano$C_{2-6}$alkenyl, $Ar^3C_{1-6}$alkyl, $(Ar^3)(OH)C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, $N(R^{8a}R^{8b})C_{1-6}$alkyl, $Ar^3C_{2-6}$alkenyl, Het-$C_{2-6}$alkenyl, $Ar^3$amino$C_{1-6}$alkyl, Het-amino$C_{1-6}$alkyl, Het-$C_{1-6}$alkylamino$C_{1-6}$alkyl, $Ar^3$thio$C_{1-6}$alkyl, Het-thio$C_{1-6}$alkyl, $Ar^3$sulfonyl$C_{1-6}$alkyl, Het-sulfonyl$C_{1-6}$alkyl, $Ar^3$aminocarbonyl, Het-aminocarbonyl, $Ar^3(CH_2)_n$aminocarbonyl, Het-$(CH_2)_n$aminocarbonyl, $Ar^3$carbonylamino, $Ar^3(CH_2)_n$amino; and the other one of $R^{2a}$ and $R^{2b}$ is hydrogen.

15. (Currently Amended) A compound according to claim 1~~any of claims 1 – 13~~, wherein one of $R^{2a}$ and $R^{3a}$ is selected from cyano$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, $N(R^{8a}R^{8b})C_{1-6}$alkyl, $Ar^3C_{2-6}$alkenyl, $Ar^3$amino$C_{1-6}$alkyl, Het-amino$C_{1-6}$alkyl, Het-$C_{1-6}$alkylamino$C_{1-6}$alkyl, $Ar^3$thio$C_{1-6}$alkyl, $Ar^3$aminocarbonyl, Het-aminocarbonyl, $Ar^3(CH_2)_n$aminocarbonyl, Het-$(CH_2)_n$aminocarbonyl; and the other one of $R^{2a}$ and $R^{2b}$ is hydrogen.

16. (Currently Amended) A compound according to claim 1~~any of claims 1 – 13~~, wherein one of $R^{2a}$ and $R^{3a}$ is selected from $N(R^{8a}R^{8b})C_{1-6}$alkyl, $Ar^3$amino$C_{1-6}$alkyl; and the other one of $R^{2a}$ and $R^{2b}$ is hydrogen.

17. (Currently Amended) A compound according to claim 1~~any of claims 14 - 16~~, wherein
in case $R^{2a}$ is hydrogen then $R^3$ is hydrogen;
in case $R^{2b}$ is hydrogen then $R^3$ is hydrogen or $C_{1-6}$alkyl.

18. (Currently Amended) A compound according to claim 1~~any of claims 1 - 17~~, wherein $R^5$ is hydrogen.

19. (Currently Amended) A compound according to claim 1~~any of claims 1 - 18~~, wherein Q is $R^{6a}$, wherein $R^{6a}$ is $C_{1-6}$alkyl substituted with one or with two substituents each independently selected from the group consisting of trifluoromethyl, $NR^{7a}R^{7b}$, $Ar^2$, hydroxy, $C_{1-4}$alkoxy, $Ar^2(CH_2)_n$oxy, hydroxycarbonyl, aminocarbonyl, $C_{1-4}$alkylcarbonyl, $C_{1-4}$alkoxycarbonyl, $Ar^2(CH_2)_n$carbonyl, aminocarbonyloxy, $C_{1-4}$alkylcarbonyloxy, $Ar^2$carbonyloxy, mono- or di($C_{1-4}$alkyl)aminocarbonyl, aminosulfonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl or a heterocycle selected from the group consisting of pyrrolidinyl, imidazolyl, piperidinyl, homopiperidinyl, piperazinyl, dioxolanyl, dioxanyl and pyridyl, wherein each of said heterocycle may optionally be substituted with with one or two radicals selected from oxo and $C_{1-6}$alkyl;

20. (Currently Amended) A compound according to claim 1~~any of claims 1 - 18~~, wherein Q is $R^{6a}$, wherein $R^{6a}$ is $C_{1-6}$alkyl substituted with $Ar^2$ or hydroxy, or $C_{1-6}$alkyl substituted with two hydroxy radicals, or $C_{1-6}$alkyl substituted with di$C_{1-6}$alkyl-dioxolanyl, pyrrolidinyl, piperidinyl, piperazinyl, 4-$C_{1-6}$alkyl-piperazinyl.

21. (Currently Amended) A compound according to claim 1~~any of claims 1 - 18~~, wherein Q is pyrrolidinyl substituted with $R^6$, piperidinyl substituted with $R^6$ or homopiperidinyl substituted with $R^6$; wherein $R^6$ is hydrogen or $C_{1-6}$alkyl optionally substituted with one or with two substituents, each independently selected from the group consisting of trifluoromethyl, $NR^{7a}R^{7b}$, $Ar^2$, hydroxy, $C_{1-4}$alkoxy, $Ar^2(CH_2)_n$oxy, hydroxycarbonyl, aminocarbonyl, $C_{1-4}$alkylcarbonyl, $C_{1-4}$alkoxycarbonyl, $Ar^2(CH_2)_n$carbonyl, aminocarbonyloxy, $C_{1-4}$alkylcarbonyloxy, $Ar^2$carbonyloxy, mono- or di($C_{1-4}$alkyl)aminocarbonyl, aminosulfonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl or a heterocycle selected from the group consisting of pyrrolidinyl, imidazolyl, piperidinyl, homopiperidinyl, piperazinyl, dioxolanyl, dioxanyl and pyridyl, wherein each of said heterocycle may optionally be substituted with with one or two radicals selected from oxo and $C_{1-6}$alkyl.

22. (Currently Amended) A compound according to claim 1~~any of claims 1 - 18~~, wherein Q is pyrrolidinyl substituted with $R^6$, piperidinyl substituted with $R^6$ or homopiperidinyl substituted with $R^6$; wherein $R^6$ is hydrogen or $C_{1-6}$alkyl optionally substituted with $NR^{7a}R^{7b}$, $Ar^2$, hydroxy, hydroxycarbonyl, aminocarbonyl, aminosulfonyl or $C_{1-6}$alkyl substituted with two hydroxy radicals, or $C_{1-6}$alkyl substituted with a heterocycle selected from dioxolanyl, pyrrolidinyl, piperidinyl, homopiperidinyl, piperazinyl, wherein each of said heterocycle may optionally be substituted with oxo or with one or two $C_{1-6}$alkyl radicals.

23. (Currently Amended) A compound according to claim 1~~any of claims 1 - 18~~, wherein Q is pyrrolidinyl substituted with $R^6$, piperidinyl substituted with $R^6$ or homopiperidinyl substituted with $R^6$; wherein $R^6$ is hydrogen or $C_{1-6}$alkyl substituted with $Ar^2$ or $C_{1-6}$alkyl substituted with piperidinyl or with piperazinyl.

24. (Currently Amended) A compound according to claim 21~~any of claims 21 - 23~~, wherein Q is piperidinyl substituted with $R^6$.

25. (Currently Amended) A compound according to claim 1~~any of claims 1 - 24~~, wherein $R^{8a}$ is $Ar^3$, $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, $C_{1-6}$alkoxy$C_{1-6}$alkyl, cyano$C_{1-6}$alkyl, amino$C_{1-6}$alkyl, mono-or di($C_{1-6}$alkyl)amino$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, aminocarbonyl-$C_{1-6}$alkyl, carboxyl-$C_{1-6}$-alkyl; and $R^{8b}$ is $Ar^3$.

26. (Currently Amended) A compound according to claim 1~~any of claims 1 - 24~~, wherein $R^{8a}$ is $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl, aminocarbonyl-$C_{1-6}$-alkyl; and $R^{8b}$ is $C_{1-6}$alkyl, hydroxy$C_{1-6}$alkyl, $Ar^3C_{1-6}$alkyl, Het-$C_{1-6}$alkyl.

27. (Currently Amended) A compound according to claim 1~~any of claims 1 - 26~~, wherein $Ar^3$ is phenyl optionally substituted with one, two or three substituents selected from halo, hydroxy, mercapto, amino, cyano, $C_{1-6}$alkyl, $C_{2-6}$alkenyl, $C_{2-6}$alkynyl, $Ar^1$, hydroxy$C_{1-6}$alkyl, $CF_3$, amino$C_{1-6}$alkyl, cyano$C_{1-6}$alkyl, aminocarbonyl, $C_{1-6}$alkyloxy, $C_{1-6}$alkylthio, $Ar^1$-oxy, $Ar^1$-thio, $Ar^1$-amino, aminosulfonyl, aminocarbonyl-$C_{1-6}$alkyl, hydroxycarbonyl-$C_{1-6}$alkyl, hydroxycarbonyl, $C_{1-4}$alkylcarbonyl, $C_{1-4}$alkylcarbonylamino or $C_{1-4}$alkoxycarbonyl.

28. (Currently Amended) A compound according to claim 1~~any of claims 1 - 27~~, wherein $Ar^3$ is phenyl substituted with one, two or three substituents selected from halo, $C_{1-6}$alkyl or hydroxy$C_{1-6}$alkyl.

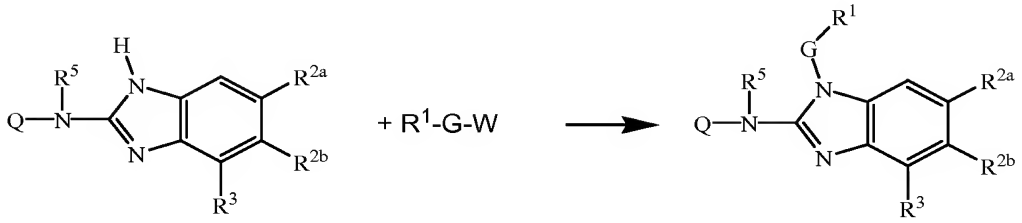
29. (Currently Amended) A compound as claimed in claim 1~~any one of claims 1 to 28~~ for use as a medicine.

30. (Currently Amended) A pharmaceutical composition comprising a pharmaceutically acceptable carrier, and as active ingredient a therapeutically effective amount of a compound as claimed in claim 1~~any one of claims 1 to 23~~.

31-32. (Cancelled)

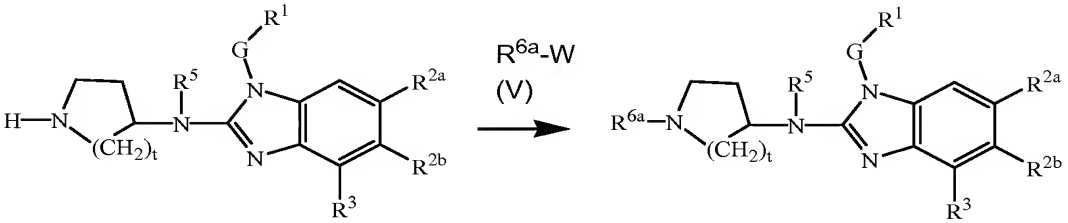
33. (Currently Amended) A process for preparing a compound as claimed in claim 1~~any of claims 1 to 23~~, said process comprising

(a) reacting an intermediate of formula (II) with a reagent (III) as in the following reaction scheme:




(II) (III) (I)

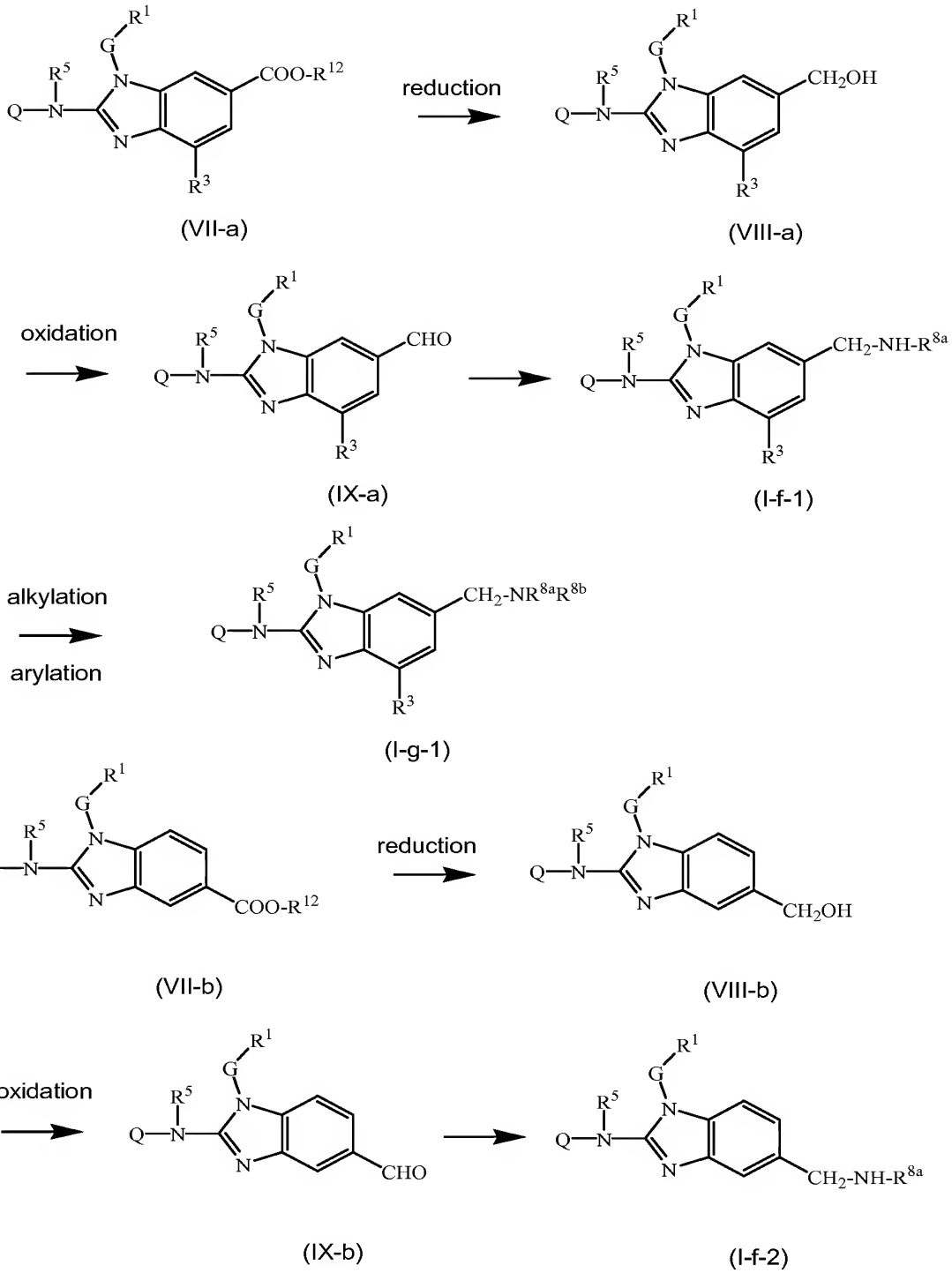
(b) reacting an intermediate of formula (IV) with a reagent (V) thus obtaining a compound of formula (I-c-1) as in the following reaction scheme:

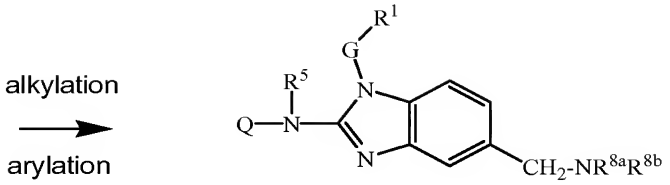



(IV) (I-c-1)

(c) reducing an intermediate (VII-a) or (VII-b) to obtain an intermediate (VIII-a) or (VIII-b) and subsequently oxidizing the alcohol group in (VIII-a) or (VIII-b) with a mild oxidant to obtain an intermediate (IX-a) or (IX-b) and subsequently alkylating (IX-a) or (IX-b) to obtain (I-f-1) or (I-f-2), which is

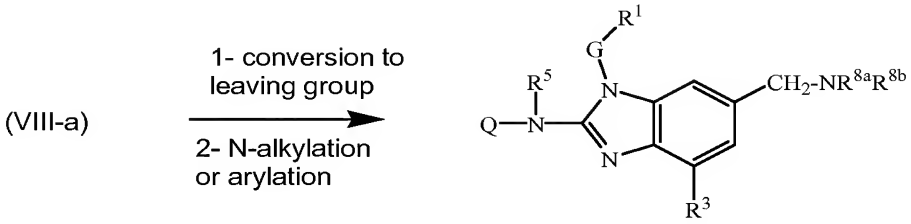
further alkylated to obtain (I-g-1) or (I-g-2) as in the following reaction schemes:



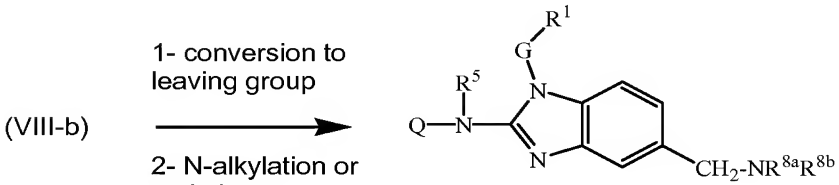



(I-g-2)

(d) converting the alcohol group in (VIII-a) or (VIII-b) to a leaving group and subsequently reacting the thus obtained products with an amine thus obtaining (I-g-1) or (I-g-2):

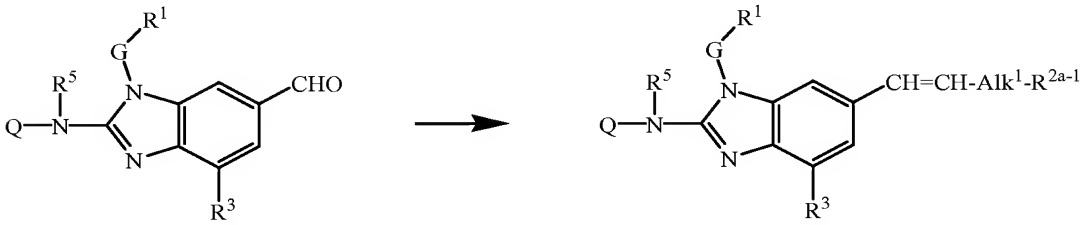



(I-g-1)

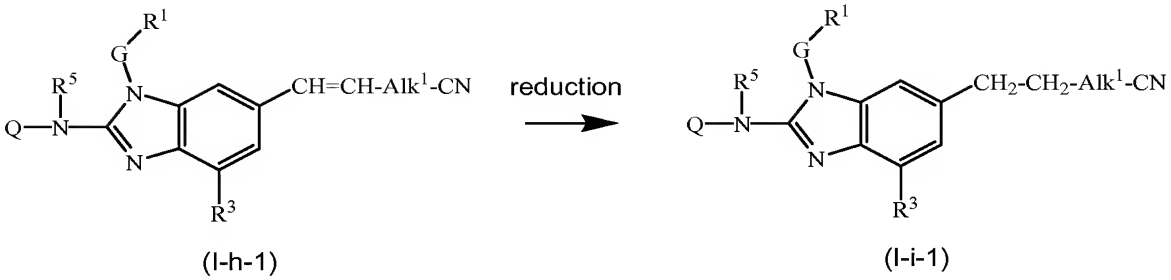



(I-g-2)

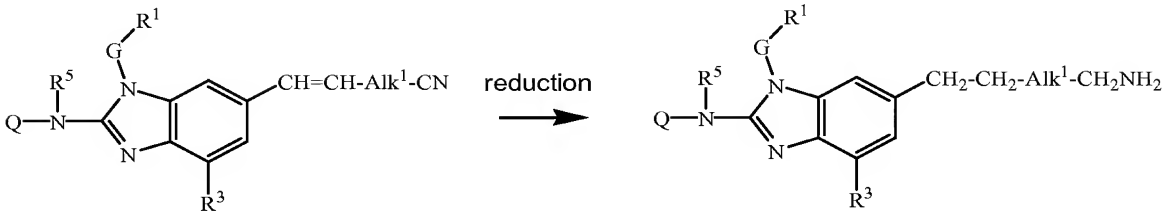
(e) converting an intermediate (IX-a) or (IX-b) to a compound (I-g-1) or (I-g-2) using a Wittig or Wittig-Horner procedure; selectively reducing the double bond in (I-g-1) or (I-g-2) thus obtaining compounds (I-i-1) or (I-i-2); reducing the cyano group in (I-i-1) or (I-i-2) to a methyleneamine group thus obtaining (I-j-1) or (I-j-2); mono- or dialkylating the latter thus obtaining compounds (I-k-1) or (I-k-2); (I-l-1) or (I-l-2):



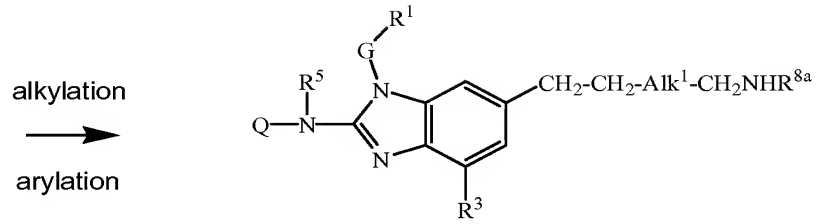

(IX-a) (I-g-1)



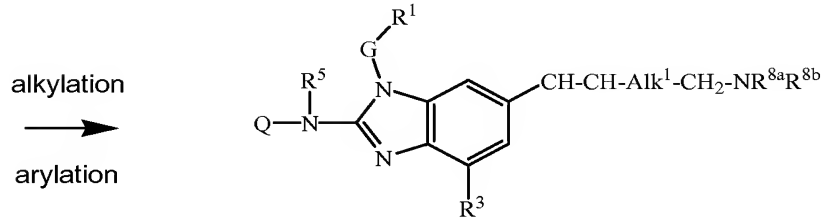

(I-h-1) (I-i-1)



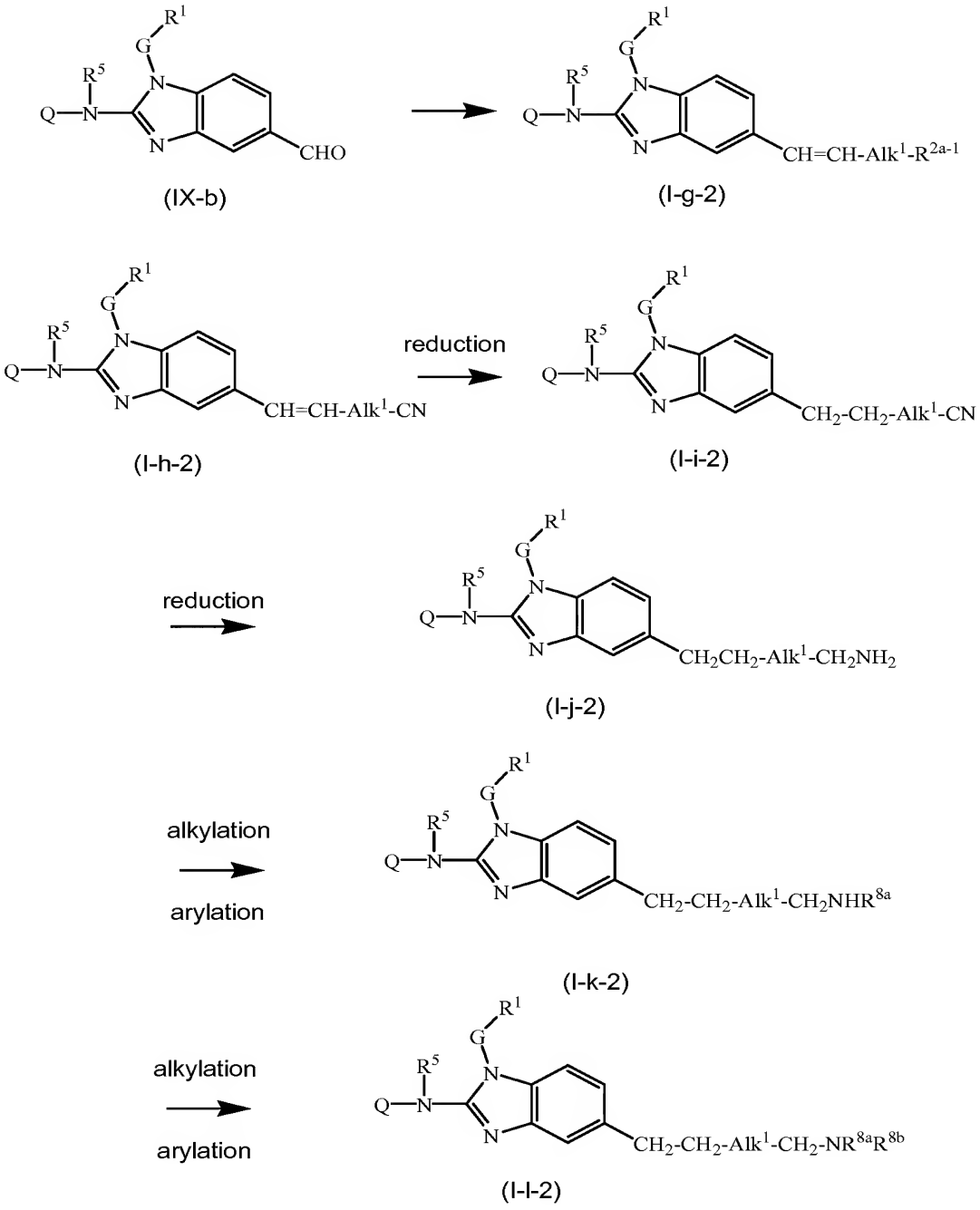

(I-h-1) (I-i-1)




(I-k-1)




(I-l-1)




and optionally converting the thus obtained compounds of formula (I) into their pharmaceutically acceptable base-addition or acid addition salt form by treatment with a suitable base or acid and conversely treating the base-addition or acid addition salt form with an acid or a base to obtain the free form of the compound of formula (I).

34. (Currently Amended) A compound of formula (VII-a), (VII-b), (VIII-a), (VIII-b), (IX-a), (IX-b), (I-f-1), (I-f-2), (I-g-1) or (I-g-2) said formula being as in claim 33, wherein G, $R^1$, $R^{2a}$, $R^{2b}$, $R^3$, $R^5$, $R^{8a}$, $R^{8b}$, $R^{12}$ are as claimed in claim 33~~claim 1~~, and wherein Q is pyrrolidinyl, piperidinyl or homopiperidinyl, substituted on their nitrogen with a radical $R^6$ which is $C_{1-6}$alkyl optionally substituted with one or two, substituents each independently selected from the group consisting of trifluoromethyl, $C_{3-7}$cycloalkyl, $Ar^2$, hydroxy, $C_{1-4}$alkoxy, $C_{1-4}$alkylthio, $Ar^2$-oxy-, $Ar^2$-thio-, $Ar^2(CH_2)_n$oxy, $Ar^2(CH_2)_n$thio, hydroxycarbonyl, aminocarbonyl, $C_{1-4}$alkylcarbonyl, $Ar^2$carbonyl, $C_{1-4}$alkoxycarbonyl, $Ar^2(CH_2)_n$carbonyl, aminocarbonyloxy, $C_{1-4}$alkylcarbonyloxy, $Ar^2$carbonyloxy, $Ar^2(CH_2)_n$carbonyloxy, $C_{1-4}$alkoxycarbonyl$(CH_2)_n$oxy, mono- or di($C_{1-4}$alkyl)aminocarbonyl, mono- or di($C_{1-4}$alkyl)aminocarbonyloxy, aminosulfonyl, mono- or di($C_{1-4}$alkyl)aminosulfonyl or a heterocycle selected from the group consisting of pyrrolidinyl, pyrrolyl, dihydropyrrolyl, imidazolyl, triazolyl, piperidinyl, homopiperidinyl, piperazinyl, dioxolanyl, dioxanyl, pyridyl and tetrahydropyridyl, wherein each of said heterocycle may optionally be substituted with one or two substituents selected from oxo or $C_{1-6}$alkyl; and wherein said $R^6$ can be represented by $R^{6b}$, as well as the pharmaceutically acceptable salt forms thereof, and the possible stereoisomeric forms thereof.

35. (Original) A compound according to claim 34 wherein $R^{6b}$ is $C_{1-6}$alkyl optionally substituted with $Ar^2$, hydroxy, aminocarbonyl, aminosulfonyl, or $C_{1-6}$alkyl substituted with two hydroxy radicals, or $C_{1-6}$alkyl substituted with pyrrolidinyl, piperidinyl, piperazinyl, 4-$C_{1-6}$alkyl-piperazinyl.

36. (Original) A compound according to claim 34 wherein $R^{6b}$ is $C_{1-6}$alkyl.

37. (Currently Amended) A compound formula (VII-a), (VII-b), (VIII-a), (VIII-b), (IX-a), (IX-b), (I-f-1), (I-f-2), (I-g-1) or (I-g-2) said formula being as in claim 33, wherein G, $R^1$, $R^{2a}$, $R^{2b}$, $R^3$, $R^5$, $R^{8a}$, $R^{8b}$ and $R^{12}$ are as claimed in claim 1 and wherein Q is $R^{6b}$ wherein $R^{6b}$ is as claimed in claim 33~~claim 1~~.

38. (Original) A compound according to claim 37 wherein $R^{6b}$ is $C_{1-6}$alkyl optionally substituted with $Ar^2$, hydroxy, aminocarbonyl, aminosulfonyl, or $C_{1-6}$alkyl substituted with two hydroxy radicals, or $C_{1-6}$alkyl substituted with pyrrolidinyl, piperidinyl, piperazinyl, 4-$C_{1-6}$alkyl-piperazinyl.

39. (Original) A compound according to claim 37 wherein $R^{6b}$ is $C_{1-6}$alkyl.

**REMARKS**

Consideration of the captioned application in view of the foregoing amendments and following remarks is requested.

The specification has been amended to refer to the priority applications.

Claims 1-30 and 33-39 are currently pending. Claims 31 and 32 are hereby cancelled and claims 7-30, 33, 34 and 37 are currently amended, without disclaimer of or prejudice to the subject matter deleted therein. No new matter has been added.

Accordingly, the claims pending and under consideration are claims 1-30 and 33-39.

Early favourable action on the merits is respectfully requested.

Applicant respectfully requests that a timely Notice of Allowance of claims 1-30 and 33-39 be issued in this case.

Respectfully submitted,

/Alana G. Kriegsman/

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